

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization
-International Bureau



(43) International Publication Date
10 July 2003 (10.07.2003)

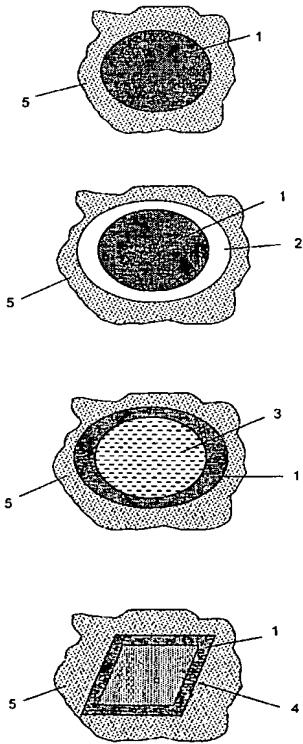
PCT

(10) International Publication Number
WO 03/055520 A1

- (51) International Patent Classification⁷: **A61K 41/00**
- (21) International Application Number: **PCT/GB02/05927**
- (22) International Filing Date:
24 December 2002 (24.12.2002)
- (25) Filing Language: English
- (26) Publication Language: English
- (30) Priority Data:
0131058.0 28 December 2001 (28.12.2001) GB
- (71) Applicant (for all designated States except US): **PSIMEI PHARMACEUTICALS PLC [GB/GB]**; 13-21 High Street, Guildford, Surrey GU1 3DG (GB).
- (72) Inventor; and
- (75) Inventor/Applicant (for US only): **PATEL, Bipin, Chandra, Muljibhai [GB/GB]**; Psimei Pharmaceuticals plc, 13-21 High Street, Guildford, Surrey GU1 3DG (GB).
- (74) Agent: **CORNISH, Kristina, Victoria, Joy; Kilburn & Strode, 20 Red Lion Street, London WC1R 4PJ (GB)**.
- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW),

[Continued on next page]

(54) Title: DELIVERY OF NEUTRON CAPTURE ELEMENTS FOR NEUTRON CAPTURE THERAPY



(57) Abstract: Neutron capture therapy (NCT) for example, Boron neutron capture therapy (BNCT) requires the delivery of a neutron capture element such as Boron to a target site to be treated, followed by irradiation with neutrons. The invention provides new means for delivery of the neutron capture element in the form of insoluble inorganic nanoparticles having a particle size of about 10^{-10} m to about 10^{-6} m. The neutron capture element can be in particulate form or in the form of glass or glass ceramic or as a polymerised inorganic matrix or as a sol-gel derived xerogel. The nanoparticles of the invention can further comprise a biocompatible outer layer which provides the function of stealth and assists in providing an appropriate clearance rate. In some embodiments, the nanoparticles comprise a core selected from, for example, mica, zeolites, TiO_2 spheres, ZrO_2 spheres or particles or organic polymer particles or spheres surrounded by a thin film of the neutron capture element. Pharmaceutical compositions, uses and methods for the treatment of cancer are disclosed. Also disclosed is a process for the preparation of water insoluble nanoparticles comprising causing friction between pure blocks of the required neutron capture element in an inorganic form and collecting nanoparticles that result therefrom.

WO 03/055520 A1

BEST AVAILABLE COPY